

This listing of claims will replace all prior versions, and listings, of claims in the application.

**Listing of Claims:**

1. (Currently amended) A method of concurrent imaging in a mammal comprising:
  - a) administering to said mammal a vitronectin receptor targeted imaging agent and a perfusion imaging agent; and
  - b) concurrently detecting the vitronectin receptor targeted imaging agent bound at the vitronectin receptor and the perfusion imaging agent; and
  - c) forming an image from the detection of said vitronectin targeted imaging agent and said perfusion imaging agent; wherein said vitronectin targeted imaging agent comprises a peptide selected from the group consisting of:

cyclo(Arg-Gly-Asp-D-Tyr(N-[[5-[carbonyl]-2-pyridinyl]diazenido]-3-aminopropyl)-Val)) SEQ ID NO: 1;

cyclo(Arg-Gly-Asp-D-Phe-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido)))  
SEQ ID NO: 3;

cyclo(Arg-Gly-Asp-D-Tyr-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido)))  
SEQ ID NO: 4;

[[5-[carbonyl]-2-pyridinyl]diazenido]-Phe-Glu(cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe} SEQ ID NO: 5;

cyclo{Arg-Gly-Asp-D-Nal-Lys([[5-[carbonyl]-2-pyridinyl]diazenido))} SEQ  
ID NO: 6

[[5-[carbonyl]-2-pyridinyl]-diazenido]-Glu(cyclo{Lys-Arg-Gly-Asp-D-Nal})-  
cyclo{Lys-Arg-Gly-Asp-D-Nal} SEQ ID NO: 7;

cyclo(Arg-Gly-Asp-D-Tyr((N-[[5-[carbonyl]-2-pyridinyl]diazenido]-18-amino-14-aza-4,7,10-oxy-15-oxo-octadecoyl)-3-aminopropyl)-Val) SEQ ID NO: 1;

N-[[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(O-cyclo(Lys-Arg-Gly-Asp-D-Phe))-O-cyclo(Lys-Arg-Gly-Asp-D-Phe) SEQ ID NO: 8;

N-[[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(O-cyclo(D-Tyr(3-aminopropyl)-Val-Arg-Gly-Asp))-O-cyclo(D-Tyr(3-aminopropyl)-Val-Arg-Gly-Asp) SEQ ID NO: 9;

cyclo(Arg-Gly-Asp-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido))-D-Val) SEQ ID NO: 10;

cyclo(N-Me-Arg-Gly-Asp-ATA-D-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido))) SEQ ID NO: 14;

cyclo(Arg-Gly-Asp-D-Tyr(N-[[5-[carbonyl]-2-pyridinyl]diazenido]-3-aminopropyl)-Val) SEQ ID NO: 1;

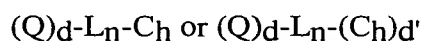
cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe} SEQ ID NO: 8;

cyclo(Arg-Gly-Asp-D-Phe-Lys) SEQ ID NO: 3; and

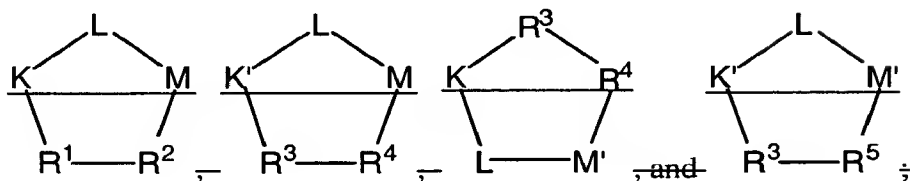
Glu-(cyclo(Arg-Gly-Asp-D-Phe-Lys)) SEQ ID NO: 16.

2. (original) The method of claim 1, wherein the vitronectin receptor is selected from the group:  $\alpha_v\beta_3$ , and  $\alpha_v\beta_5$ .

3. (original) The method according to claim 1, wherein the vitronectin receptor is  $\alpha_v\beta_3$ .
4. (original) The method of claim 1 wherein the perfusion imaging agent is selected from the group consisting of: an ultrasound perfusion agent, an MRI perfusion imaging agent, and a radiolabelled imaging agent.
5. (original) The method of claim 1 wherein the perfusion imaging agent is hexakis methoxyisobutyl isonitrile Technetium(I) ( $^{99m}\text{Tc}$ -Sestamibi),  $^{210}\text{Tl}$ ,  $^{99m}\text{Tc}$ -tetrofosmin,  $^{99m}\text{Tc}$ -furifosmin, or  $^{99m}\text{Tc}$ -NOET.
6. (original) The method according to claim 1, wherein the vitronectin receptor targeted imaging agent is a diagnostic metallopharmaceutical.
7. (original) The method according to claim 6, wherein the vitronectin receptor targeting agent is a vitronectin antagonist.
8. (original) The method according to claim 6, wherein the vitronectin receptor targeting agent is a vitronectin agonist.
9. (original) The method of claim 6, wherein the diagnostic metallopharmaceutical comprises a metal and a compound, wherein the compound comprises:
  - a) a chelator capable of chelating the metal;
  - b) a targeting moiety, wherein the targeting moiety is bound to the chelator; and
  - c) 0-1 linking groups between the targeting moiety and the chelator;wherein the targeting moiety is a peptide or peptidomimetic which binds to a vitronectin receptor.
10. (Currently amended) The method according to claim 9, wherein compound is of the formula:



wherein, Q is a peptide independently selected from the group:



~~K is an L-amino acid independently selected at each occurrence from the group: arginine, citrulline, N-methylarginine, lysine, homolysine, 2-aminoethyleysteine,  $\delta$ -N-2-imidazolinylnornithine,  $\delta$ -N-benzylcarbameoylnornithine, and  $\beta$ -2-benzimidazolylacetyl-1,2-diaminopropionic acid;~~

~~K' is a D-amino acid independently selected at each occurrence from the group: arginine, citrulline, N-methylarginine, lysine, homolysine, 2-aminoethyleysteine,  $\delta$ -N-2-imidazolinylnornithine,  $\delta$ -N-benzylcarbameoylnornithine, and  $\beta$ -2-benzimidazolylacetyl-1,2-diaminopropionic acid;~~

~~L is independently selected at each occurrence from the group: glycine, L-alanine, and D-alanine;~~

~~M is L-aspartic acid;~~

~~M' is D-aspartic acid;~~

~~R<sup>1</sup> is an amino acid substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each occurrence from the group: glycine, L-valine, D-valine, alanine, leucine, isoleucine, norleucine, 2-aminobutyric acid, 2-aminohexanoic acid, tyrosine, phenylalanine, thienylalanine, phenylglycine, cyclohexylalanine, homophenylalanine,~~

~~1-naphthylalanine, lysine, serine, ornithine, 1,2-diaminobutyric acid,  
1,2-diaminopropionic acid, cysteine, penicillamine, and methionine;~~

~~R<sup>2</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, valine, alanine, leucine, isoleucine, norleucine,  
2-aminobutyric acid, 2-aminohexanoic acid, tyrosine, L-phenylalanine, D-  
phenylalanine, thienylalanine, phenylglycine, biphenylglycine, cyclohexylalanine,  
homophenylalanine, L-1-naphthylalanine, D-1-naphthylalanine, lysine, serine,  
ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic acid, cysteine,  
penicillamine, methionine, and 2-aminothiazole-4-acetic acid;~~

~~R<sup>3</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, D-valine, D-alanine, D-leucine, D-isoleucine,  
D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic acid, D-tyrosine,  
D-phenylalanine, D-thienylalanine, D-phenylglycine, D-cyclohexylalanine,  
D-homophenylalanine, D-1-naphthylalanine, D-lysine, D-serine, D-ornithine,  
D-1,2-diaminobutyric acid, D-1,2-diaminopropionic acid, D-cysteine,  
D-penicillamine, and D-methionine;~~

~~R<sup>4</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, D-valine, D-alanine, D-leucine, D-isoleucine,  
D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic acid, D-tyrosine,  
D-phenylalanine, D-thienylalanine, D-phenylglycine, D-cyclohexylalanine,  
D-homophenylalanine, D-1-naphthylalanine, D-lysine, D-serine, D-ornithine,  
D-1,2-diaminobutyric acid, D-1,2-diaminopropionic acid, D-cysteine,  
D-penicillamine, D-methionine, and 2-aminothiazole-4-acetic acid;~~

~~R<sup>5</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, L-valine, L-alanine, L-leucine, L-isoleucine,~~

~~L-norleucine, L-2-aminobutyric acid, L-2-aminohexanoic acid, L-tyrosine,  
L-phenylalanine, L-thienylalanine, L-phenylglycine, L-cyclohexylalanine,  
L-homophenylalanine, L-1-naphthylalanine, L-lysine, L-serine, L-ornithine,  
L-1,2-diaminobutyric acid, L-1,2-diaminopropionic acid, L-cysteine, L-penicillamine,  
L-methionine, and 2-aminothiazole-4-acetic acid;~~

~~provided that one of R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, and R<sup>5</sup> in each Q is substituted with a bond to L<sub>n</sub>;~~

~~further provided that when R<sup>2</sup> is 2-aminothiazole-4-acetic acid, K is~~

~~N-methylarginine, further provided that when R<sup>4</sup> is 2-aminothiazole-4-acetic acid, K~~

~~and K' are N-methylarginine, and still further provided that when R<sup>5</sup> is~~

~~2-aminothiazole-4-acetic acid, K' is N-methylarginine;~~

~~d is selected from 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10 selected from the group consisting of:~~

cyclo(Arg-Gly-Asp-D-Tyr(N-[[5-[carbonyl]-2-pyridinyl]diazenido]-3-aminopropyl)-Val)) SEQ ID NO: 1;

cyclo(Arg-Gly-Asp-D-Phe-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido)))  
SEQ ID NO: 3;

cyclo(Arg-Gly-Asp-D-Tyr-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido)))  
SEQ ID NO: 4;

[[5-[carbonyl]-2-pyridinyl]diazenido]-Phe-Glu(cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe} SEQ ID NO: 5;

cyclo{Arg-Gly-Asp-D-Nal-Lys([[5-[carbonyl]-2-pyridinyl]diazenido))} SEQ  
ID NO: 6

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PATENT

[[5-[carbonyl]-2-pyridinyl]-diazenido]-Glu(cyclo{Lys-Arg-Gly-Asp-D-Nal})-  
cyclo{Lys-Arg-Gly-Asp-D-Nal} SEQ ID NO: 7;

cyclo(Arg-Gly-Asp-D-Tyr((N-[[5-[carbonyl]-2-pyridinyl]diazenido]-18-  
amino-14-aza-4,7,10-oxy-15-oxo-octadecoyl)-3-aminopropyl)-Val) SEQ ID  
NO: 1;

N-[[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(O-cyclo(Lys-Arg-Gly-Asp-D-  
Phe))-O-cyclo(Lys-Arg-Gly-Asp-D-Phe) SEQ ID NO: 8;

N-[[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(O-cyclo(D-Tyr(3-aminopropyl)-  
Val-Arg-Gly-Asp))-O-cyclo(D-Tyr(3-aminopropyl)-Val-Arg-Gly-Asp) SEQ  
ID NO: 9;

cyclo(Arg-Gly-Asp-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido))-D-Val)  
SEQ ID NO: 10;

cyclo(N-Me-Arg-Gly-Asp-ATA-D-Lys(N-[[5-[carbonyl]-2-  
pyridinyl]diazenido))) SEQ ID NO: 14;

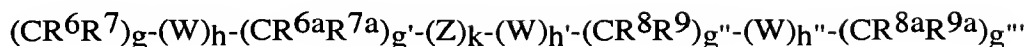
cyclo(Arg-Gly-Asp-D-Tyr(N-[[5-[carbonyl]-2-pyridinyl]diazenido]-3-  
aminopropyl)-Val) SEQ ID NO: 1;

cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe} SEQ ID  
NO: 8;

cyclo(Arg-Gly-Asp-D-Phe-Lys) SEQ ID NO: 3; and

Glu-(cyclo(Arg-Gly-Asp-D-Phe-Lys)) SEQ ID NO: 16;

$L_n$  is a linking group having the formula:



provided that  $g+h+g'+k+h'+g''+h''+g'''$  is other than 0;

W is independently selected at each occurrence from the group: O, S, NH, NHC(=O), C(=O)NH, C(=O), C(=O)O, OC(=O), NHC(=S)NH, NHC(=O)NH, SO<sub>2</sub>, (OCH<sub>2</sub>CH<sub>2</sub>)<sub>s</sub>, (CH<sub>2</sub>CH<sub>2</sub>O)<sub>s'</sub>, (OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>)<sub>s''</sub>, (CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>O)<sub>t</sub>, and (aa)<sub>t'</sub>;

aa is independently at each occurrence an amino acid;

Z is selected from the group: aryl substituted with 0-3 R<sup>10</sup>, C<sub>3-10</sub> cycloalkyl substituted with 0-3 R<sup>10</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>10</sup>;

R<sup>6</sup>, R<sup>6a</sup>, R<sup>7</sup>, R<sup>7a</sup>, R<sup>8</sup>, R<sup>8a</sup>, R<sup>9</sup> and R<sup>9a</sup> are independently selected at each occurrence from the group: H, =O, COOH, SO<sub>3</sub>H, PO<sub>3</sub>H, C<sub>1-5</sub> alkyl substituted with 0-3 R<sup>10</sup>, aryl substituted with 0-3 R<sup>10</sup>, benzyl substituted with 0-3 R<sup>10</sup>, and C<sub>1-5</sub> alkoxy substituted with 0-3 R<sup>10</sup>, NHC(=O)R<sup>11</sup>, C(=O)NHR<sup>11</sup>, NHC(=O)NHR<sup>11</sup>, NHR<sup>11</sup>, R<sup>11</sup>, and a bond to C<sub>H</sub>;

R<sup>10</sup> is independently selected at each occurrence from the group: a bond to C<sub>H</sub>, COOR<sup>11</sup>, OH, NHR<sup>11</sup>, SO<sub>3</sub>H, PO<sub>3</sub>H, aryl substituted with 0-3 R<sup>11</sup>, C<sub>1-5</sub> alkyl substituted with 0-1 R<sup>12</sup>, C<sub>1-5</sub> alkoxy substituted with 0-1 R<sup>12</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>11</sup>;



$R^{11}$  is independently selected at each occurrence from the group: H, aryl substituted with 0-1  $R^{12}$ , a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-1  $R^{12}$ ,  $C_{3-10}$  cycloalkyl substituted with 0-1  $R^{12}$ , polyalkylene glycol substituted with 0-1  $R^{12}$ , carbohydrate substituted with 0-1  $R^{12}$ , cyclodextrin substituted with 0-1  $R^{12}$ , amino acid substituted with 0-1  $R^{12}$ , polycarboxyalkyl substituted with 0-1  $R^{12}$ , polyazaalkyl substituted with 0-1  $R^{12}$ , peptide substituted with 0-1  $R^{12}$ , wherein the peptide is comprised of 2-10 amino acids, and a bond to  $C_H$ ;

$R^{12}$  is a bond to  $C_H$ ;

k is selected from 0, 1, and 2;

h is selected from 0, 1, and 2;

h' is selected from 0, 1, 2, 3, 4, and 5;

h'' is selected from 0, 1, 2, 3, 4, and 5;

g is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g'' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g''' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

s is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

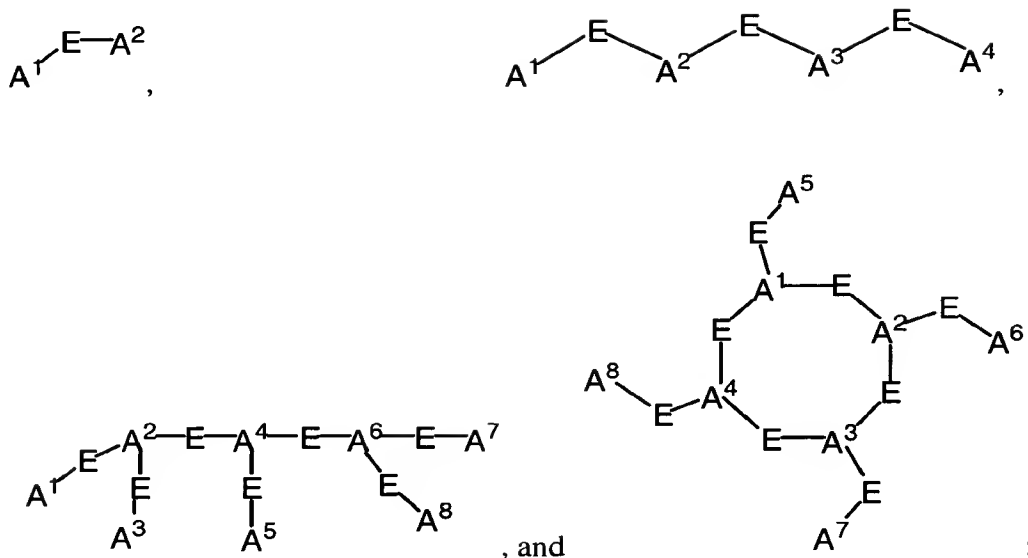
s' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

s'' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

t is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

t' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

$C_H$  is a metal bonding unit having a formula selected from the group:



A<sup>1</sup>, A<sup>2</sup>, A<sup>3</sup>, A<sup>4</sup>, A<sup>5</sup>, A<sup>6</sup>, A<sup>7</sup>, and A<sup>8</sup> are independently selected at each occurrence from the group N, NR<sup>13</sup>, NR<sup>13</sup>R<sup>14</sup>, S, SH, O, OH, PR<sup>13</sup>, PR<sup>13</sup>R<sup>14</sup>, P(O)R<sup>15</sup>R<sup>16</sup>, and a bond to L<sub>n</sub>;

E is a bond, CH, or a spacer group independently selected at each occurrence from the group:

C<sub>1</sub>-C<sub>10</sub> alkyl substituted with 0-3 R<sup>17</sup>, aryl substituted with 0-3 R<sup>17</sup>, C<sub>3</sub>-<sub>10</sub> cycloalkyl substituted with 0-3 R<sup>17</sup>, heterocyclo-C<sub>1</sub>-<sub>10</sub> alkyl substituted with 0-3 R<sup>17</sup>, wherein the heterocyclo group is a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O, C<sub>6</sub>-<sub>10</sub> aryl-C<sub>1</sub>-<sub>10</sub> alkyl substituted with 0-3 R<sup>17</sup>, C<sub>1</sub>-<sub>10</sub> alkyl-C<sub>6</sub>-<sub>10</sub> aryl- substituted with 0-3 R<sup>17</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>17</sup>;

R<sup>13</sup>, and R<sup>14</sup> are each independently selected from the group: a bond to L<sub>n</sub>, hydrogen,

C<sub>1</sub>-C<sub>10</sub> alkyl substituted with 0-3 R<sup>17</sup>, aryl substituted with 0-3 R<sup>17</sup>, C<sub>1</sub>-<sub>10</sub> cycloalkyl substituted with 0-3 R<sup>17</sup>, heterocyclo-C<sub>1</sub>-<sub>10</sub> alkyl substituted with 0-3

R<sup>17</sup>, wherein the heterocyclo group is a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O, C<sub>6-10</sub> aryl-C<sub>1-10</sub> alkyl substituted with 0-3 R<sup>17</sup>, C<sub>1-10</sub> alkyl-C<sub>6-10</sub> aryl- substituted with 0-3 R<sup>17</sup>, a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>17</sup>, and an electron, provided that when one of R<sup>13</sup> or R<sup>14</sup> is an electron, then the other is also an electron;

alternatively, R<sup>13</sup> and R<sup>14</sup> combine to form =C(R<sup>20</sup>)(R<sup>21</sup>);

R<sup>15</sup> and R<sup>16</sup> are each independently selected from the group: a bond to L<sub>n</sub>, -OH, C<sub>1-C10</sub> alkyl substituted with 0-3 R<sup>17</sup>, C<sub>1-C10</sub> alkyl substituted with 0-3 R<sup>17</sup>, aryl substituted with 0-3 R<sup>17</sup>, C<sub>3-10</sub> cycloalkyl substituted with 0-3 R<sup>17</sup>, heterocyclo-C<sub>1-10</sub> alkyl substituted with 0-3 R<sup>17</sup>, wherein the heterocyclo group is a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O, C<sub>6-10</sub> aryl-C<sub>1-10</sub> alkyl substituted with 0-3 R<sup>17</sup>, C<sub>1-10</sub> alkyl-C<sub>6-10</sub> aryl- substituted with 0-3 R<sup>17</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>17</sup>;

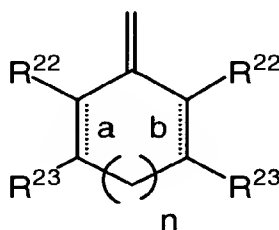
R<sup>17</sup> is independently selected at each occurrence from the group: a bond to L<sub>n</sub>, =O, F, Cl, Br, I, -CF<sub>3</sub>, -CN, -CO<sub>2</sub>R<sup>18</sup>, -C(=O)R<sup>18</sup>, -C(=O)N(R<sup>18</sup>)<sub>2</sub>, -CHO, -CH<sub>2</sub>OR<sup>18</sup>, -OC(=O)R<sup>18</sup>, -OC(=O)OR<sup>18a</sup>, -OR<sup>18</sup>, -OC(=O)N(R<sup>18</sup>)<sub>2</sub>, -NR<sup>19</sup>C(=O)R<sup>18</sup>, -NR<sup>19</sup>C(=O)OR<sup>18a</sup>, -NR<sup>19</sup>C(=O)N(R<sup>18</sup>)<sub>2</sub>, -NR<sup>19</sup>SO<sub>2</sub>N(R<sup>18</sup>)<sub>2</sub>, -NR<sup>19</sup>SO<sub>2</sub>R<sup>18a</sup>, -SO<sub>3</sub>H, -SO<sub>2</sub>R<sup>18a</sup>, -SR<sup>18</sup>, -S(=O)R<sup>18a</sup>, -SO<sub>2</sub>N(R<sup>18</sup>)<sub>2</sub>, -N(R<sup>18</sup>)<sub>2</sub>, -NHC(=S)NHR<sup>18</sup>, =NOR<sup>18</sup>, NO<sub>2</sub>, -C(=O)NHOR<sup>18</sup>, -C(=O)NHN(R<sup>18</sup>)R<sup>18a</sup>,

-OCH<sub>2</sub>CO<sub>2</sub>H, 2-(1-morpholino)ethoxy, C<sub>1</sub>-C<sub>5</sub> alkyl, C<sub>2</sub>-C<sub>4</sub> alkenyl, C<sub>3</sub>-C<sub>6</sub> cycloalkyl, C<sub>3</sub>-C<sub>6</sub> cycloalkylmethyl, C<sub>2</sub>-C<sub>6</sub> alkoxyalkyl, aryl substituted with 0-2 R<sup>18</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O;

R<sup>18</sup>, R<sup>18a</sup>, and R<sup>19</sup> are independently selected at each occurrence from the group: a bond to L<sub>n</sub>, H, C<sub>1</sub>-C<sub>6</sub> alkyl, phenyl, benzyl, C<sub>1</sub>-C<sub>6</sub> alkoxy, halide, nitro, cyano, and trifluoromethyl;

R<sup>20</sup> and R<sup>21</sup> are independently selected from the group: H, C<sub>1</sub>-C<sub>10</sub> alkyl, -CN, -CO<sub>2</sub>R<sup>25</sup>, -C(=O)R<sup>25</sup>, -C(=O)N(R<sup>25</sup>)<sub>2</sub>, C<sub>2</sub>-C<sub>10</sub> 1-alkene substituted with 0-3 R<sup>23</sup>, C<sub>2</sub>-C<sub>10</sub> 1-alkyne substituted with 0-3 R<sup>23</sup>, aryl substituted with 0-3 R<sup>23</sup>, unsaturated 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>23</sup>, and unsaturated C<sub>3</sub>-10 carbocycle substituted with 0-3 R<sup>23</sup>;

alternatively, R<sup>20</sup> and R<sup>21</sup>, taken together with the divalent carbon radical to which they are attached form:



R<sup>22</sup> and R<sup>23</sup> are independently selected from the group: H, R<sup>24</sup>, C<sub>1</sub>-C<sub>10</sub> alkyl substituted with 0-3 R<sup>24</sup>, C<sub>2</sub>-C<sub>10</sub> alkenyl substituted with 0-3 R<sup>24</sup>, C<sub>2</sub>-C<sub>10</sub> alkynyl substituted with 0-3 R<sup>24</sup>, aryl substituted with 0-3 R<sup>24</sup>, a 5-10 membered heterocyclic ring

system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>24</sup>, and C<sub>3-10</sub> carbocycle substituted with 0-3 R<sup>24</sup>;

alternatively, R<sup>22</sup>, R<sup>23</sup> taken together form a fused aromatic or a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O;

a and b indicate the positions of optional double bonds and n is 0 or 1;

R<sup>24</sup> is independently selected at each occurrence from the group: =O, F, Cl, Br, I, -CF<sub>3</sub>, -CN, -CO<sub>2</sub>R<sup>25</sup>, -C(=O)R<sup>25</sup>, -C(=O)N(R<sup>25</sup>)<sub>2</sub>, -N(R<sup>25</sup>)<sub>3</sub><sup>+</sup>, -CH<sub>2</sub>OR<sup>25</sup>, -OC(=O)R<sup>25</sup>, -OC(=O)OR<sup>25a</sup>, -OR<sup>25</sup>, -OC(=O)N(R<sup>25</sup>)<sub>2</sub>, -NR<sup>26</sup>C(=O)R<sup>25</sup>, -NR<sup>26</sup>C(=O)OR<sup>25a</sup>, -NR<sup>26</sup>C(=O)N(R<sup>25</sup>)<sub>2</sub>, -NR<sup>26</sup>SO<sub>2</sub>N(R<sup>25</sup>)<sub>2</sub>, -NR<sup>26</sup>SO<sub>2</sub>R<sup>25a</sup>, -SO<sub>3</sub>H, -SO<sub>2</sub>R<sup>25a</sup>, -SR<sup>25</sup>, -S(=O)R<sup>25a</sup>, -SO<sub>2</sub>N(R<sup>25</sup>)<sub>2</sub>, -N(R<sup>25</sup>)<sub>2</sub>, =NOR<sup>25</sup>, -C(=O)NHOR<sup>25</sup>, -OCH<sub>2</sub>CO<sub>2</sub>H, and 2-(1-morpholino)ethoxy; and,

R<sup>25</sup>, R<sup>25a</sup>, and R<sup>26</sup> are each independently selected at each occurrence from the group: hydrogen and C<sub>1</sub>-C<sub>6</sub> alkyl;

and a pharmaceutically acceptable salt thereof.

11. (Currently amended) The method according to claim 10 wherein

L is glycine;

~~R<sup>1</sup> is an amino acid, optionally substituted with a bond to L<sub>n</sub>, independently selected at each occurrence from the group: L-valine, D-valine, alanine, leucine, isoleucine, norleucine, 2-aminobutyric acid, tyrosine, phenylalanine, phenylglycine, cyclohexylalanine, homophenylalanine, lysine, ornithine, 1,2-diaminobutyric acid, and 1,2-diaminopropionic acid;~~

~~R<sup>2</sup> is an amino acid, optionally substituted with a bond to L<sub>n</sub>, independently selected at each occurrence from the group: valine, alanine, leucine, isoleucine, norleucine, 2-aminobutyric acid, tyrosine, L-phenylalanine, D-phenylalanine, thienylalanine, phenylglycine, biphenylglycine, cyclohexylalanine, homophenylalanine, L-1-naphthylalanine, D-1-naphthylalanine, lysine, ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic acid, and 2-aminothiazole-4-acetic acid;~~

~~R<sup>3</sup> is an amino acid, optionally substituted with a bond to L<sub>n</sub>, independently selected at each occurrence from the group: D-valine, D-alanine, D-leucine, D-isoleucine, D-norleucine, D-2-aminobutyric acid, D-tyrosine, D-phenylalanine, D-phenylglycine, D-cyclohexylalanine, D-homophenylalanine, D-lysine, D-serine, D-ornithine, D-1,2-diaminobutyric acid, and D-1,2-diaminopropionic acid;~~

~~R<sup>4</sup> is an amino acid, optionally substituted with a bond to L<sub>n</sub>, independently selected at each occurrence from the group: D-valine, D-alanine, D-leucine, D-isoleucine, D-norleucine, D-2-aminobutyric acid, D-tyrosine, D-phenylalanine, D-thienylalanine, D-phenylglycine, D-cyclohexylalanine, D-homophenylalanine, D-1-naphthylalanine, D-lysine, D-ornithine, D-1,2-diaminobutyric acid, D-1,2-diaminopropionic acid, and 2-aminothiazole-4-acetic acid;~~

~~R<sup>5</sup> is an amino acid, optionally substituted with a bond to L<sub>n</sub>, independently selected at each occurrence from the group: L-valine, L-alanine, L-leucine, L-isoleucine, L-norleucine, L-2-aminobutyric acid, L-tyrosine, L-phenylalanine, L-thienylalanine, L-phenylglycine, L-cyclohexylalanine, L-homophenylalanine, L-1-naphthylalanine, L-lysine, L-ornithine, L-1,2-diaminobutyric acid, L-1,2-diaminopropionic acid, and 2-aminothiazole-4-acetic acid;~~

d is selected from 1, 2, and 3;

W is independently selected at each occurrence from the group: O, NH, NHC(=O), C(=O)NH, C(=O), C(=O)O, OC(=O), NHC(=S)NH, NHC(=O)NH, SO<sub>2</sub>, (OCH<sub>2</sub>CH<sub>2</sub>)<sub>s</sub>, (CH<sub>2</sub>CH<sub>2</sub>O)<sub>s'</sub>, (OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>)<sub>s''</sub>, and (CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>O)<sub>t</sub>,

Z is selected from the group: aryl substituted with 0-1 R<sup>10</sup>, C<sub>3-10</sub> cycloalkyl substituted with 0-1 R<sup>10</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-1 R<sup>10</sup>; R<sup>6</sup>, R<sup>6a</sup>, R<sup>7</sup>, R<sup>7a</sup>, R<sup>8</sup>, R<sup>8a</sup>, R<sup>9</sup>, and R<sup>9a</sup> are independently selected at each occurrence from the group: H, =O, COOH, SO<sub>3</sub>H, C<sub>1-5</sub> alkyl substituted with 0-1 R<sup>10</sup>, aryl substituted with 0-1 R<sup>10</sup>, benzyl substituted with 0-1 R<sup>10</sup>, and C<sub>1-5</sub> alkoxy substituted with 0-1 R<sup>10</sup>, NHC(=O)R<sup>11</sup>, C(=O)NHR<sup>11</sup>, NHC(=O)NHR<sup>11</sup>, NHR<sup>11</sup>, R<sup>11</sup>, and a bond to C<sub>H</sub>;

R<sup>10</sup> is independently selected at each occurrence from the group: COOR<sup>11</sup>, OH, NHR<sup>11</sup>, SO<sub>3</sub>H, aryl substituted with 0-1 R<sup>11</sup>, a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-1 R<sup>11</sup>, C<sub>1-5</sub> alkyl substituted with 0-1 R<sup>12</sup>, C<sub>1-5</sub> alkoxy substituted with 0-1 R<sup>12</sup>, and a bond to C<sub>H</sub>;

R<sup>11</sup> is independently selected at each occurrence from the group: H, aryl substituted with 0-1 R<sup>12</sup>, a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-1 R<sup>12</sup>, polyalkylene glycol substituted with 0-1 R<sup>12</sup>, carbohydrate substituted with 0-1 R<sup>12</sup>, cyclodextrin substituted with 0-1 R<sup>12</sup>, amino acid substituted with 0-1 R<sup>12</sup>, and a bond to C<sub>H</sub>;

k is 0 or 1;

h is 0 or 1;

h' is 0 or 1;

s is selected from 0, 1, 2, 3, 4, and 5;

s' is selected from 0, 1, 2, 3, 4, and 5;

s" is selected from 0, 1, 2, 3, 4, and 5;

t is selected from 0, 1, 2, 3, 4, and 5;

A<sup>1</sup>, A<sup>2</sup>, A<sup>3</sup>, A<sup>4</sup>, A<sup>5</sup>, A<sup>6</sup>, A<sup>7</sup>, and A<sup>8</sup> are independently selected at each occurrence from the group: NR<sup>13</sup>, NR<sup>13</sup>R<sup>14</sup>, S, SH, OH, and a bond to L<sub>n</sub>;

E is a bond, CH, or a spacer group independently selected at each occurrence from the group:

C<sub>1</sub>-C<sub>10</sub> alkyl substituted with 0-3 R<sup>17</sup>, aryl substituted with 0-3 R<sup>17</sup>, C<sub>3</sub>-<sub>10</sub> cycloalkyl substituted with 0-3 R<sup>17</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>17</sup>;

R<sup>13</sup>, and R<sup>14</sup> are each independently selected from the group: a bond to L<sub>n</sub>, hydrogen, C<sub>1</sub>-C<sub>10</sub> alkyl substituted with 0-3 R<sup>17</sup>, aryl substituted with 0-3 R<sup>17</sup>, a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>17</sup>, and an electron, provided that when one of R<sup>13</sup> or R<sup>14</sup> is an electron, then the other is also an electron;

alternatively, R<sup>13</sup> and R<sup>14</sup> combine to form =C(R<sup>20</sup>)(R<sup>21</sup>);

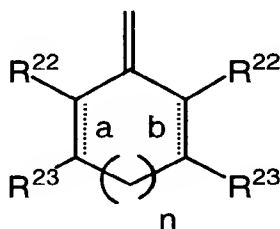
R<sup>17</sup> is independently selected at each occurrence from the group: a bond to L<sub>n</sub>, =O, F, Cl, Br, I, -CF<sub>3</sub>, -CN, -CO<sub>2</sub>R<sup>18</sup>, -C(=O)R<sup>18</sup>, -C(=O)N(R<sup>18</sup>)<sub>2</sub>, -CH<sub>2</sub>OR<sup>18</sup>, -OC(=O)R<sup>18</sup>, -OC(=O)OR<sup>18a</sup>, -OR<sup>18</sup>, -OC(=O)N(R<sup>18</sup>)<sub>2</sub>, -NR<sup>19</sup>C(=O)R<sup>18</sup>, -NR<sup>19</sup>C(=O)OR<sup>18a</sup>, -NR<sup>19</sup>C(=O)N(R<sup>18</sup>)<sub>2</sub>, -NR<sup>19</sup>SO<sub>2</sub>N(R<sup>18</sup>)<sub>2</sub>, -NR<sup>19</sup>SO<sub>2</sub>R<sup>18a</sup>, -SO<sub>3</sub>H, -SO<sub>2</sub>R<sup>18a</sup>, -S(=O)R<sup>18a</sup>, -SO<sub>2</sub>N(R<sup>18</sup>)<sub>2</sub>, -N(R<sup>18</sup>)<sub>2</sub>, -NHC(=S)NHR<sup>18</sup>, =NOR<sup>18</sup>, -C(=O)NHN(R<sup>18</sup>)R<sup>18a</sup>, -OCH<sub>2</sub>CO<sub>2</sub>H, and 2-(1-morpholino)ethoxy;

R<sup>18</sup>, R<sup>18a</sup>, and R<sup>19</sup> are independently selected at each occurrence from the group: a bond to L<sub>n</sub>, H, and C<sub>1</sub>-C<sub>6</sub> alkyl;



$R^{20}$  and  $R^{21}$  are independently selected from the group: H, C<sub>1</sub>-C<sub>5</sub> alkyl, -CO<sub>2</sub> $R^{25}$ , C<sub>2</sub>-C<sub>5</sub> 1-alkene substituted with 0-3  $R^{23}$ , C<sub>2</sub>-C<sub>5</sub> 1-alkyne substituted with 0-3  $R^{23}$ , aryl substituted with 0-3  $R^{23}$ , and unsaturated 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3  $R^{23}$ ;

alternatively,  $R^{20}$  and  $R^{21}$ , taken together with the divalent carbon radical to which they are attached form:



$R^{22}$  and  $R^{23}$  are independently selected from the group: H, and  $R^{24}$ ;

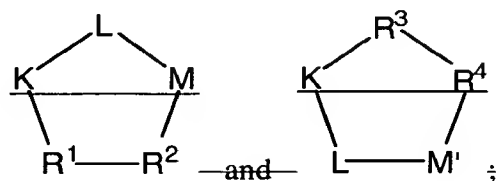
alternatively,  $R^{22}$ ,  $R^{23}$  taken together form a fused aromatic or a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O;

$R^{24}$  is independently selected at each occurrence from the group: -CO<sub>2</sub> $R^{25}$ , -C(=O)N( $R^{25}$ )<sub>2</sub>, -CH<sub>2</sub>OR<sup>25</sup>, -OC(=O) $R^{25}$ , -OR<sup>25</sup>, -SO<sub>3</sub>H, -N( $R^{25}$ )<sub>2</sub>, and -OCH<sub>2</sub>CO<sub>2</sub>H; and,

$R^{25}$  is independently selected at each occurrence from the group: H and C<sub>1</sub>-C<sub>3</sub> alkyl.

12. (Currently amended) The method according to claim 10 wherein

~~Q is a peptide selected from the group:~~



~~R<sup>1</sup> is L-valine, D-valine, D-lysine optionally substituted on the ε amino group with a bond to L<sub>n</sub> or L-lysine optionally substituted on the ε amino group with a bond to L<sub>n</sub>;~~

~~R<sup>2</sup> is L-phenylalanine, D-phenylalanine, D-1-naphthylalanine, 2-aminothiazole-4-acetic acid, L-lysine optionally substituted on the ε amino group with a bond to L<sub>n</sub> or tyrosine, the tyrosine optionally substituted on the hydroxy group with a bond to L<sub>n</sub>;~~

~~R<sup>3</sup> is D-valine, D-phenylalanine, or L-lysine optionally substituted on the ε amino group with a bond to L<sub>n</sub>;~~

~~R<sup>4</sup> is D-phenylalanine, D-tyrosine substituted on the hydroxy group with a bond to L<sub>n</sub>, or L-lysine optionally substituted on the ε amino group with a bond to L<sub>n</sub>;~~

~~provided that one of R<sup>1</sup> and R<sup>2</sup> in each Q is substituted with a bond to L<sub>n</sub>, and further~~

~~provided that when R<sup>2</sup> is 2-aminothiazole-4-acetic acid, K is N-methylarginine;~~

~~d is 1 or 2;~~

W is independently selected at each occurrence from the group:  $\text{NHC(=O)}$ ,  $\text{C(=O)NH}$ ,  $\text{C(=O)}$ ,  $(\text{CH}_2\text{CH}_2\text{O})_{s'}$ , and  $(\text{CH}_2\text{CH}_2\text{CH}_2\text{O})_t$ ;

$\text{R}^6$ ,  $\text{R}^{6a}$ ,  $\text{R}^7$ ,  $\text{R}^{7a}$ ,  $\text{R}^8$ ,  $\text{R}^{8a}$ ,  $\text{R}^9$ , and  $\text{R}^{9a}$  are independently selected at each occurrence from the group: H,  $\text{NHC(=O)R}^{11}$ , and a bond to  $\text{C}_h$ ;

k is 0;

h" is selected from 0, 1, 2, and 3;

g is selected from 0, 1, 2, 3, 4, and 5;

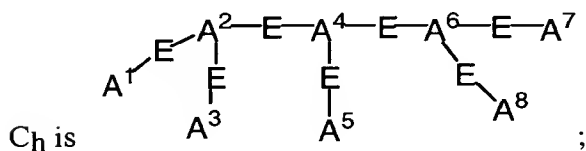
g' is selected from 0, 1, 2, 3, 4, and 5;

g" is selected from 0, 1, 2, 3, 4, and 5;

g''' is selected from 0, 1, 2, 3, 4, and 5;

s' is 1 or 2;

t is 1 or 2;



$\text{A}^1$  is selected from the group: OH, and a bond to  $\text{L}_n$ ;

$\text{A}^2$ ,  $\text{A}^4$ , and  $\text{A}^6$  are each N;

$\text{A}^3$ ,  $\text{A}^5$ , and  $\text{A}^8$  are each OH;

$\text{A}^7$  is a bond to  $\text{L}_n$  or NH-bond to  $\text{L}_n$ ;

E is a  $\text{C}_2$  alkyl substituted with 0-1  $\text{R}^{17}$ ;

$R^{17}$  is =O;

alternatively,  $C_h$  is  $A^1 \begin{array}{c} \diagup \\ E-A^2 \end{array}$ ;

$A^1$  is  $NH_2$  or  $N=C(R^{20})(R^{21})$ ;

E is a bond;

$A^2$  is  $NHR^{13}$ ;

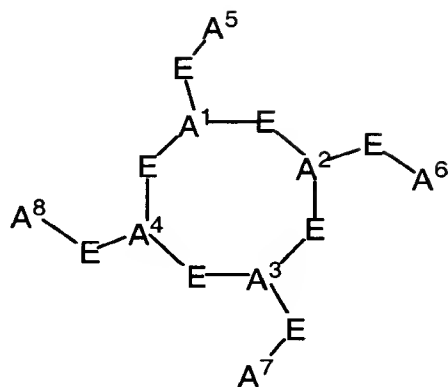
$R^{13}$  is a heterocycle substituted with  $R^{17}$ , the heterocycle being selected from pyridine and pyrimidine;

$R^{17}$  is selected from a bond to  $L_n$ ,  $C(=O)NHR^{18}$ , and  $C(=O)R^{18}$ ;

$R^{18}$  is a bond to  $L_n$ ;

$R^{24}$  is selected from the group:  $-CO_2R^{25}$ ,  $-OR^{25}$ ,  $-SO_3H$ , and  $-N(R^{25})_2$ ;

$R^{25}$  is independently selected at each occurrence from the group: hydrogen and methyl;



alternatively,  $C_h$  is

$A^1$ ,  $A^2$ ,  $A^3$ , and  $A^4$  are each N;

$A^5$ ,  $A^6$ , and  $A^8$  are each OH;

$A^7$  is a bond to  $L_n$ ;

E is a  $C_2$  alkyl substituted with 0-1  $R^{17}$ ; and,

$R^{17}$  is =O.

13. (original) The method of claim 6 wherein the diagnostic metallopharmaceutical comprises a radioisotope.

14. (original) The method of claim 13 wherein the radioisotope is selected from the group consisting of  $^{99m}\text{Tc}$ ,  $^{95}\text{Tc}$ ,  $^{111}\text{In}$ ,  $^{62}\text{Cu}$ ,  $^{64}\text{Cu}$ ,  $^{67}\text{Ga}$ , and  $^{68}\text{Ga}$ .

15. (original) The method of claim 14 wherein the radioisotope is selected from the group consisting of In-111, and Tc-99m.

16. (original) The method of claim 9, wherein the metallopharmaceutical is a diagnostic radiopharmaceutical and the metal is a radioisotope selected from the group:  $^{99m}\text{Tc}$ ,  $^{95}\text{Tc}$ ,  $^{111}\text{In}$ ,  $^{62}\text{Cu}$ ,  $^{64}\text{Cu}$ ,  $^{67}\text{Ga}$ , and  $^{68}\text{Ga}$ .

17. (original) The method of claim 16 wherein the radioisotope is selected from the group consisting of  $^{111}\text{In}$ , and  $^{99m}\text{Tc}$ .

18. (original) The method according to claim 16, wherein the radioisotope is  $^{99m}\text{Tc}$  or  $^{95}\text{Tc}$ , the radiopharmaceutical further comprises a first ancillary ligand and a second ancillary ligand capable of stabilizing the radiopharmaceutical.

19. (original) The method according to claim 16, wherein the radioisotope is  $^{99m}\text{Tc}$ .

20. (currently amended) The method according to claim 19, wherein the radiopharmaceutical is selected from the group:

$^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}(\text{Arg-Gly-Asp-D-Tyr}(\text{N}-[[5\text{-[carbonyl]-2-pyridinyl}] \text{diazenido}] \text{-3-aminopropyl})\text{-Val}))$  SEQ ID NO: 1;

~~$^{99m}\text{Tc}(\text{tricine})(\text{TPPMS})(\text{cyclo}(\text{Arg-D-Val-D-Tyr}(\text{N}-[[5\text{-[carbonyl]-2-pyridinyl}] \text{diazenido}] \text{-3-aminopropyl})\text{-D-Asp-Gly}))$  SEQ ID NO: 2;~~

~~$^{99m}\text{Tc}(\text{tricine})(\text{TPPDS})(\text{cyclo}(\text{Arg-D-Val-D-Tyr}(\text{N}-[[5\text{-[carbonyl]-2-pyridinyl}] \text{diazenido}] \text{-3-aminopropyl})\text{-D-Asp-Gly}))$  SEQ ID NO: 2;~~

~~$^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}(\text{Arg-D-Val-D-Tyr}(\text{N}-[[5\text{-[carbonyl]-2-pyridinyl}] \text{diazenido}] \text{-3-aminopropyl})\text{-D-Asp-Gly}))$  SEQ ID NO: 2;~~

$^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}(\text{Arg-Gly-Asp-D-Phe-Lys}(\text{N}-[[5\text{-[carbonyl]-2-pyridinyl}] \text{diazenido}] \text{))))}$  SEQ ID NO: 3;

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**Application No.:** 09/995,388  
**Office Action Dated:** March 15, 2005

**PATENT**

<sup>99m</sup>Tc(tricine)(TPPTS)(cyclo(Arg-Gly-Asp-D-Tyr-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido)))) SEQ ID NO: 4;

<sup>99m</sup>Tc(tricine)(TPPTS)([[5-[carbonyl]-2-pyridinyl]diazenido]-Phe-Glu(cyclo{Lys-Arg-Gly-Asp-D-Phe}))-cyclo{Lys-Arg-Gly-Asp-D-Phe}) SEQ ID NO: 5;

<sup>99m</sup>Tc(tricine)(TPPTS)(cyclo{ Arg-Gly-Asp-D-Nal-Lys([[5-[carbonyl]-2-pyridinyl]diazenido)]}) SEQ ID NO: 6;

<sup>99m</sup>Tc(tricine)(TPPTS)([[5-[carbonyl]-2-pyridinyl]-diazenido]-Glu(cyclo{Lys-Arg-Gly-Asp-D-Nal}))-cyclo{Lys-Arg-Gly-Asp-D-Nal}) SEQ ID NO: 7;

<sup>99m</sup>Tc(tricine)(TPPTS)(cyclo(Arg-Gly-Asp-D-Tyr((N-[[5-[carbonyl]-2-pyridinyl]diazenido]-18-amino-14-aza-4,7,10-oxy-15-oxo-octadecoyl)-3-aminopropyl)-Val)) SEQ ID NO: 1;

<sup>99m</sup>Tc(tricine)(TPPTS)(N-[[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(O-cyclo(Lys-Arg-Gly-Asp-D-Phe)))-O-cyclo(Lys-Arg-Gly-Asp-D-Phe)) SEQ ID NO: 8;

<sup>99m</sup>Tc(tricine)(TPPTS)(N-[[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(O-cyclo(D-Tyr(3-aminopropyl)-Val-Arg-Gly-Asp)))-O-cyclo(D-Tyr(3-aminopropyl)-Val-Arg-Gly-Asp)) SEQ ID NO: 9;

<sup>99m</sup>Tc(tricine)(TPPTS)(cyclo(Arg-Gly-Asp-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido))-D-Val)) SEQ ID NO: 10;

~~<sup>99m</sup>Tc(tricine)(TPPTS)(cyclo{D-Lys([[5-[carbonyl]-2-pyridinyl]diazenido))-D-Phe-D-Asp-Gly-Arg}) SEQ ID NO: 11;~~

~~<sup>99m</sup>Tc(tricine)(TPPTS)([[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(cyclo{D-Lys-D-Phe-D-Asp-Gly-Arg}))-cyclo{D-Lys-D-Phe-D-Asp-Gly-Arg}) SEQ ID NO: 12;~~

~~<sup>99m</sup>Tc(tricine)(TPPTS)(cyclo{D-Phe-D-Lys([5-[carbonyl]-2-pyridinyl]diazenido))-D-Asp-Gly-Arg}) SEQ ID NO: 13;~~

<sup>99m</sup>Tc(tricine)(TPPTS)(cyclo(N-Me-Arg-Gly-Asp-ATA-D-Lys(N-[5-[carbonyl]-2-pyridinyl]diazenido)))) SEQ ID NO: 14;

~~<sup>99m</sup>Tc(tricine)(TPPTS)(cyclo{Cit-Gly-Asp-D-Phe-Lys([5-[carbonyl]-2-pyridinyl]diazenido))}) SEQ ID NO: 15; and~~

<sup>99m</sup>Tc(tricine)(1,2,4-triazole)(cyclo(Arg-Gly-Asp-D-Tyr(N-[5-[carbonyl]-2-pyridinyl]diazenido)-3-aminopropyl)-Val)) SEQ ID NO: 1.

21. (original) The method according to claim 16, wherein the radioisotope is <sup>111</sup>In.

22. (Previously amended) The method according to claim 21, wherein the radiopharmaceutical is selected from the group:

(DOTA-<sup>111</sup>In)-Glu(cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe} SEQ ID NO: 8;

cyclo(Arg-Gly-Asp-D-Phe-Lys(DTPA-<sup>111</sup>In)) SEQ ID NO: 3; and,

cyclo(Arg-Gly-Asp-D-Phe-Lys)<sub>2</sub>(DTPA-<sup>111</sup>In) SEQ ID NO: 3.

23. (original) The method according to claim 6 wherein the diagnostic metallopharmaceutical is comprised of a paramagnetic metal.

24. (original) The method according to claim 23 wherein the paramagnetic metal is selected from the group consisting of Gd(III), Dy(III), Fe(III) and Mn(II).



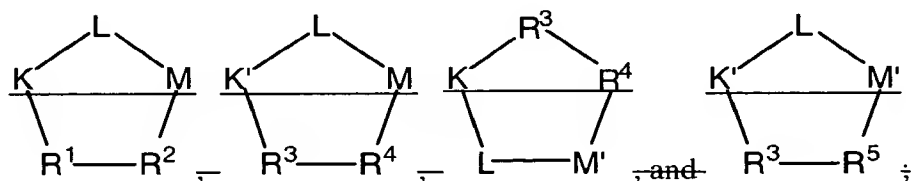
25. (original) The method according to claim 23 wherein the paramagnetic metal is Gd(III).
26. (original) The method according to claim 9, wherein the metal is a paramagnetic metal ion selected from the group Gd(III), Dy(III), Fe(III) and Mn(II).
27. (original) The method according to claim 26, wherein the metal ion is Gd(III).
28. (Previously amended) The method according to claim 27, wherein the contrast agent is:  
cyclo(Arg-Gly-Asp-D-Tyr(N-DTPA(Gd(III))-3-aminopropyl)-Val) SEQ ID NO: 1.
29. (original) The method according to claim 6 wherein the diagnostic metallopharmaceutical is a X-ray contrast agent.
30. (original) The method according to claim 29 wherein the X-ray contrast agent comprises a vitronectin targeting agent; and the metal is selected from the group: Re, Sm, Ho, Lu, Pm, Y, Bi, Pd, Gd, La, Au, Au, Yb, Dy, Cu, Rh, Ag, and Ir.
31. (original) The method according to claim 9, wherein diagnostic metallopharmaceutical is a X-ray contrast agent; the metal is selected from the group: Re, Sm, Ho, Lu, Pm, Y, Bi, Pd, Gd, La, Au, Au, Yb, Dy, Cu, Rh, Ag, and Ir.
32. (original) A kit comprising a compound of claim 9 and a perfusion imaging agent.
33. (original) The kit of Claim 32 further comprising a reducing agent.
34. (original) The kit of Claim 33 wherein the reducing agent is tin(II).
35. (original) The kit of Claim 33 further comprising one or more ancillary ligands.

36. (original) The kit of Claim 35 wherein the ancillary ligands are tricine and TPPTS.
37. (original) A kit comprising a compound of claim 10 and a perfusion imaging agent.
38. (original) A method according to claim 1, wherein the vitronectin targeted imaging agent is a vitronectin targeted ultrasound imaging agent.
39. (original) A method according to Claim 38, wherein the ultrasound imaging agent comprises an echogenic gas or temperature activated gaseous precursor, and a compound, wherein the compound comprises:
- a) a surfactant;
  - b) a targeting moiety, wherein the targeting moiety is bound to the surfactant; and
  - c) 0-1 linking groups between the targeting moiety and surfactant;
- wherein the targeting moiety is a peptide or peptidomimetic, which binds to a vitronectin receptor.

40. (Currently amended) A method according to Claim 39, wherein the compound is of the formula:



wherein, Q is a cyclic pentapeptide ~~independently selected from the group:~~



~~K is an L-amino acid independently selected at each occurrence from the group: arginine, citrulline, N-methylarginine, lysine, homolysine, 2-aminoethylcysteine,~~

~~$\delta$ -N-2-imidazolinylnornithine,  $\delta$ -N-benzylcarbamoylnornithine, and  
 $\beta$ -2-benzimidazolylacetyl-1,2-diaminopropionic acid;~~

~~K' is a D-amino acid independently selected at each occurrence from the group: arginine,  
citrulline, N-methylarginine, lysine, homolysine, 2-aminoethyleysteine,  
 $\delta$ -N-2-imidazolinylnornithine,  $\delta$ -N-benzylcarbamoylnornithine, and  
 $\beta$ -2-benzimidazolylacetyl-1,2-diaminopropionic acid;~~

~~L is independently selected at each occurrence from the group: glycine, L-alanine, and  
D-alanine;~~

~~M is L-aspartic acid;~~

~~M' is D-aspartic acid;~~

~~R<sup>1</sup> is an amino acid substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, L-valine, D-valine, alanine, leucine, isoleucine,  
norleucine, 2-aminobutyric acid, 2-aminohexanoic acid, tyrosine, phenylalanine,  
thienylalanine, phenylglycine, cyclohexylalanine, homophenylalanine,  
1-naphthylalanine, lysine, serine, ornithine, 1,2-diaminobutyric acid,  
1,2-diaminopropionic acid, cysteine, penicillamine, and methionine;~~

~~R<sup>2</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, valine, alanine, leucine, isoleucine, norleucine,  
2-aminobutyric acid, 2-aminohexanoic acid, tyrosine, L-phenylalanine, D-  
phenylalanine, thienylalanine, phenylglycine, biphenylglycine, cyclohexylalanine,  
homophenylalanine, L-1-naphthylalanine, D-1-naphthylalanine, lysine, serine,  
ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic acid, cysteine,  
penicillamine, methionine, and 2-aminothiazole-4-acetic acid;~~

~~R<sup>3</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each occurrence from the group: glycine, D-valine, D-alanine, D-leucine, D-isoleucine, D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic acid, D-tyrosine, D-phenylalanine, D-thienylalanine, D-phenylglycine, D-cyclohexylalanine, D-homophenylalanine, D-1-naphthylalanine, D-lysine, D-serine, D-ornithine, D-1,2-diaminobutyric acid, D-1,2-diaminopropionic acid, D-cysteine, D-penicillamine, and D-methionine;~~

~~R<sup>4</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each occurrence from the group: glycine, D-valine, D-alanine, D-leucine, D-isoleucine, D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic acid, D-tyrosine, D-phenylalanine, D-thienylalanine, D-phenylglycine, D-cyclohexylalanine, D-homophenylalanine, D-1-naphthylalanine, D-lysine, D-serine, D-ornithine, D-1,2-diaminobutyric acid, D-1,2-diaminopropionic acid, D-cysteine, D-penicillamine, D-methionine, and 2-aminothiazole-4-acetic acid;~~

~~R<sup>5</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each occurrence from the group: glycine, L-valine, L-alanine, L-leucine, L-isoleucine, L-norleucine, L-2-aminobutyric acid, L-2-aminohexanoic acid, L-tyrosine, L-phenylalanine, L-thienylalanine, L-phenylglycine, L-cyclohexylalanine, L-homophenylalanine, L-1-naphthylalanine, L-lysine, L-serine, L-ornithine, L-1,2-diaminobutyric acid, L-1,2-diaminopropionic acid, L-cysteine, L-penicillamine, L-methionine, and 2-aminothiazole-4-acetic acid;~~

~~provided that one of R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, and R<sup>5</sup> in each Q is substituted with a bond to L<sub>n</sub>;~~

~~further provided that when R<sup>2</sup> is 2-aminothiazole-4-acetic acid, K is~~

~~N-methylarginine, further provided that when R<sup>4</sup> is 2-aminothiazole-4-acetic acid, K~~

and K' are N-methylarginine, and still further provided that when R<sup>5</sup> is  
2-aminothiazole-4-acetic acid, K' is N-methylarginine;

d is selected from 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10 selected from the group consisting of:

cyclo(Arg-Gly-Asp-D-Tyr(N-[[5-[carbonyl]-2-pyridinyl]diazenido]-3-aminopropyl)-Val)) SEQ ID NO: 1;

cyclo(Arg-Gly-Asp-D-Phe-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido)))  
SEQ ID NO: 3;

cyclo(Arg-Gly-Asp-D-Tyr-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido)))  
SEQ ID NO: 4;

[[5-[carbonyl]-2-pyridinyl]diazenido]-Phe-Glu(cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe} SEQ ID NO: 5;

cyclo{Arg-Gly-Asp-D-Nal-Lys([5-[carbonyl]-2-pyridinyl]diazenido))} SEQ  
ID NO: 6

[[5-[carbonyl]-2-pyridinyl]-diazenido]-Glu(cyclo{Lys-Arg-Gly-Asp-D-Nal))-  
cyclo{Lys-Arg-Gly-Asp-D-Nal} SEQ ID NO: 7;

cyclo(Arg-Gly-Asp-D-Tyr((N-[[5-[carbonyl]-2-pyridinyl]diazenido]-18-  
amino-14-aza-4,7,10-oxy-15-oxo-octadecoyl)-3-aminopropyl)-Val) SEQ ID  
NO: 1;

N-[[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(O-cyclo(Lys-Arg-Gly-Asp-D-  
Phe))-O-cyclo(Lys-Arg-Gly-Asp-D-Phe) SEQ ID NO: 8;

N-[[5-[carbonyl]-2-pyridinyl]diazenido]-Glu(O-cyclo(D-Tyr(3-aminopropyl)-Val-Arg-Gly-Asp))-O-cyclo(D-Tyr(3-aminopropyl)-Val-Arg-Gly-Asp) SEQ ID NO: 9;

cyclo(Arg-Gly-Asp-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido))-D-Val) SEQ ID NO: 10;

cyclo(N-Me-Arg-Gly-Asp-ATA-D-Lys(N-[[5-[carbonyl]-2-pyridinyl]diazenido))) SEQ ID NO: 14;

cyclo(Arg-Gly-Asp-D-Tyr(N-[[5-[carbonyl]-2-pyridinyl]diazenido]-3-aminopropyl)-Val) SEQ ID NO: 1;

cyclo{Lys-Arg-Gly-Asp-D-Phe))-cyclo{Lys-Arg-Gly-Asp-D-Phe} SEQ ID NO: 8;

cyclo(Arg-Gly-Asp-D-Phe-Lys) SEQ ID NO: 3; and

Glu-(cyclo(Arg-Gly-Asp-D-Phe-Lys)) SEQ ID NO: 16;

S<sub>f</sub> is a surfactant which is a lipid or a compound of the formula:  $A^9-E^1-A^{10}$  ;

A<sup>9</sup> is selected from the group: OH and OR<sup>27</sup>;

A<sup>10</sup> is OR<sup>27</sup>;

R<sup>27</sup> is C(=O)C<sub>1-20</sub> alkyl;

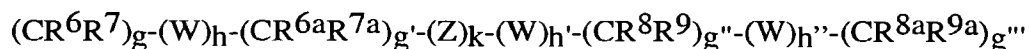
E<sup>1</sup> is C<sub>1-10</sub> alkylene substituted with 1-3 R<sup>28</sup>;

R<sup>28</sup> is independently selected at each occurrence from the group: R<sup>30</sup>, -PO<sub>3</sub>H-R<sup>30</sup>, =O, -CO<sub>2</sub>R<sup>29</sup>, -C(=O)R<sup>29</sup>, -C(=O)N(R<sup>29</sup>)<sub>2</sub>, -CH<sub>2</sub>OR<sup>29</sup>, -OR<sup>29</sup>, -N(R<sup>29</sup>)<sub>2</sub>, C<sub>1</sub>-C<sub>5</sub> alkyl, and C<sub>2</sub>-C<sub>4</sub> alkenyl;

R<sup>29</sup> is independently selected at each occurrence from the group: R<sup>30</sup>, H, C<sub>1</sub>-C<sub>6</sub> alkyl, phenyl, benzyl, and trifluoromethyl;

R<sup>30</sup> is a bond to L<sub>n</sub>;

L<sub>n</sub> is a linking group having the formula:



W is independently selected at each occurrence from the group: O, S, NH, NHC(=O), C(=O)NH, C(=O), C(=O)O, OC(=O), NHC(=S)NH, NHC(=O)NH, SO<sub>2</sub>, (OCH<sub>2</sub>CH<sub>2</sub>)<sub>20-200</sub>, (CH<sub>2</sub>CH<sub>2</sub>O)<sub>20-200</sub>, (OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>)<sub>20-200</sub>, (CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>O)<sub>20-200</sub>, and (aa)<sub>t</sub>;

aa is independently at each occurrence an amino acid;

Z is selected from the group: aryl substituted with 0-3 R<sup>10</sup>, C<sub>3</sub>-10 cycloalkyl substituted with 0-3 R<sup>10</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>10</sup>;

R<sup>6</sup>, R<sup>6a</sup>, R<sup>7</sup>, R<sup>7a</sup>, R<sup>8</sup>, R<sup>8a</sup>, R<sup>9</sup> and R<sup>9a</sup> are independently selected at each occurrence from the group: H, =O, COOH, SO<sub>3</sub>H, PO<sub>3</sub>H, C<sub>1</sub>-C<sub>5</sub> alkyl substituted with 0-3 R<sup>10</sup>, aryl substituted with 0-3 R<sup>10</sup>, benzyl substituted with 0-3 R<sup>10</sup>, and C<sub>1</sub>-C<sub>5</sub> alkoxy

substituted with 0-3  $R^{10}$ ,  $NHC(=O)R^{11}$ ,  $C(=O)NHR^{11}$ ,  $NHC(=O)NHR^{11}$ ,  $NHR^{11}$ ,  $R^{11}$ , and a bond to  $S_f$ ;

$R^{10}$  is independently selected at each occurrence from the group: a bond to  $S_f$ ,  $COOR^{11}$ ,  $OH$ ,  $NHR^{11}$ ,  $SO_3H$ ,  $PO_3H$ , aryl substituted with 0-3  $R^{11}$ ,  $C_{1-5}$  alkyl substituted with 0-1  $R^{12}$ ,  $C_{1-5}$  alkoxy substituted with 0-1  $R^{12}$ , and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3  $R^{11}$ ;

$R^{11}$  is independently selected at each occurrence from the group: H, aryl substituted with 0-1  $R^{12}$ , a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-1  $R^{12}$ ,  $C_{3-10}$  cycloalkyl substituted with 0-1  $R^{12}$ , amino acid substituted with 0-1  $R^{12}$ , and a bond to  $S_f$ ;

$R^{12}$  is a bond to  $S_f$ ;

k is selected from 0, 1, and 2;

h is selected from 0, 1, and 2;

h' is selected from 0, 1, 2, 3, 4, and 5;

h'' is selected from 0, 1, 2, 3, 4, and 5;

g is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g'' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g''' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

t' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

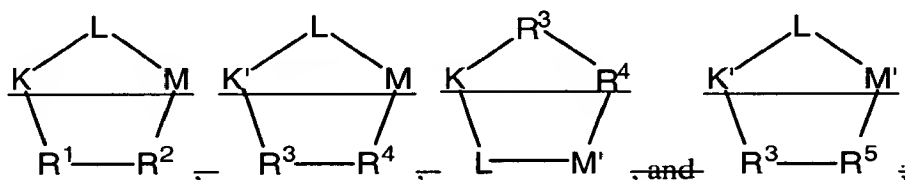
and a pharmaceutically acceptable salt thereof.



41. (Currently amended) A method according to Claim 40, wherein ~~the compound is of~~  
 the formula:



wherein, ~~Q is a cyclic pentapeptide independently selected from the group:~~



~~N-methylarginine, lysine, homolysine, 2-aminoethylcysteine,  $\delta$ -N-2-imidazolinylnornithine,  
 $\delta$ -N-benzylcarbamoylnornithine, and  $\beta$ -2-benzimidazolylacetyl-1,2-diaminopropionic  
 acid;~~

~~K' is a D-amino acid independently selected at each occurrence from the group: arginine,  
 citrulline, N-methylarginine, lysine, homolysine, 2-aminoethylcysteine,  
 $\delta$ -N-2-imidazolinylnornithine,  $\delta$ -N-benzylcarbamoylnornithine, and  
 $\beta$ -2-benzimidazolylacetyl-1,2-diaminopropionic acid;~~

~~L is independently selected at each occurrence from the group: glycine, L-alanine, and  
 D-alanine;~~

~~M is L-aspartic acid;~~

~~M' is D-aspartic acid;~~

~~R<sup>1</sup> is an amino acid substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
 occurrence from the group: glycine, L-valine, D-valine, alanine, leucine, isoleucine,  
 norleucine, 2-aminobutyric acid, 2-aminohexanoic acid, tyrosine, phenylalanine,~~

~~thienylalanine, phenylglycine, cyclohexylalanine, homophenylalanine,  
1-naphthylalanine, lysine, serine, ornithine, 1,2-diaminobutyric acid,  
1,2-diaminopropionic acid, cysteine, penicillamine, and methionine;~~

~~R<sup>2</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, valine, alanine, leucine, isoleucine, norleucine,  
2-aminobutyric acid, 2-aminohexanoic acid, tyrosine, L-phenylalanine, D-  
phenylalanine, thienylalanine, phenylglycine, biphenylglycine, cyclohexylalanine,  
homophenylalanine, L-1-naphthylalanine, D-1-naphthylalanine, lysine, serine,  
ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic acid, cysteine,  
penicillamine, methionine, and 2-aminothiazole-4-acetic acid;~~

~~R<sup>3</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, D-valine, D-alanine, D-leucine, D-isoleucine,  
D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic acid, D-tyrosine,  
D-phenylalanine, D-thienylalanine, D-phenylglycine, D-cyclohexylalanine,  
D-homophenylalanine, D-1-naphthylalanine, D-lysine, D-serine, D-ornithine,  
D-1,2-diaminobutyric acid, D-1,2-diaminopropionic acid, D-cysteine,  
D-penicillamine, and D-methionine;~~

~~R<sup>4</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each  
occurrence from the group: glycine, D-valine, D-alanine, D-leucine, D-isoleucine,  
D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic acid, D-tyrosine,  
D-phenylalanine, D-thienylalanine, D-phenylglycine, D-cyclohexylalanine,  
D-homophenylalanine, D-1-naphthylalanine, D-lysine, D-serine, D-ornithine,  
D-1,2-diaminobutyric acid, D-1,2-diaminopropionic acid, D-cysteine,  
D-penicillamine, D-methionine, and 2-aminothiazole-4-acetic acid;~~

~~R<sup>5</sup> is an amino acid, substituted with 0-1 bonds to L<sub>n</sub>, independently selected at each occurrence from the group: glycine, L-valine, L-alanine, L-leucine, L-isoleucine, L-norleucine, L-2-aminobutyric acid, L-2-aminohexanoic acid, L-tyrosine, L-phenylalanine, L-thienylalanine, L-phenylglycine, L-cyclohexylalanine, L-homophenylalanine, L-1-naphthylalanine, L-lysine, L-serine, L-ornithine, L-1,2-diaminobutyric acid, L-1,2-diaminopropionic acid, L-cysteine, L-penicillamine, L-methionine, and 2-aminothiazole-4-acetic acid;~~

~~provided that one of R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, and R<sup>5</sup> in each Q is substituted with a bond to L<sub>n</sub>;  
further provided that when R<sup>2</sup> is 2-aminothiazole-4-acetic acid, K is N-methylarginine, further provided that when R<sup>4</sup> is 2-aminothiazole-4-acetic acid, K and K' are N-methylarginine, and still further provided that when R<sup>5</sup> is 2-aminothiazole-4-acetic acid, K' is N-methylarginine;~~

S<sub>f</sub> is a surfactant which is a lipid or a compound of the formula:  $A^9-E^1-A^{10}$  ;

A<sup>9</sup> is OR<sup>27</sup>;

A<sup>10</sup> is OR<sup>27</sup>;

R<sup>27</sup> is C(=O)C<sub>1-15</sub> alkyl;

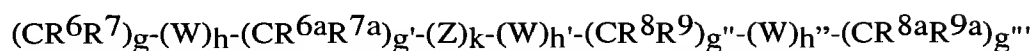
E<sup>1</sup> is C<sub>1-4</sub> alkylene substituted with 1-3 R<sup>28</sup>;

R<sup>28</sup> is independently selected at each occurrence from the group: R<sup>30</sup>, -PO<sub>3</sub>H-R<sup>30</sup>, =O, -CO<sub>2</sub>R<sup>29</sup>, -C(=O)R<sup>29</sup>, -CH<sub>2</sub>OR<sup>29</sup>, -OR<sup>29</sup>, and C<sub>1</sub>-C<sub>5</sub> alkyl;

R<sup>29</sup> is independently selected at each occurrence from the group: R<sup>30</sup>, H, C<sub>1</sub>-C<sub>6</sub> alkyl, phenyl, and benzyl;

R<sup>30</sup> is a bond to L<sub>n</sub>;

L<sub>n</sub> is a linking group having the formula:



W is independently selected at each occurrence from the group: O, S, NH, NHC(=O), C(=O)NH, C(=O), C(=O)O, OC(=O), NHC(=S)NH, NHC(=O)NH, SO<sub>2</sub>, (OCH<sub>2</sub>CH<sub>2</sub>)<sub>20-200</sub>, (CH<sub>2</sub>CH<sub>2</sub>O)<sub>20-200</sub>, (OCH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>)<sub>20-200</sub>, (CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>O)<sub>20-200</sub>, and (aa)<sub>t</sub>;

aa is independently at each occurrence an amino acid;

Z is selected from the group: aryl substituted with 0-3 R<sup>10</sup>, C<sub>3-10</sub> cycloalkyl substituted with 0-3 R<sup>10</sup>, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R<sup>10</sup>;

R<sup>6</sup>, R<sup>6a</sup>, R<sup>7</sup>, R<sup>7a</sup>, R<sup>8</sup>, R<sup>8a</sup>, R<sup>9</sup> and R<sup>9a</sup> are independently selected at each occurrence from the group: H, =O, C<sub>1</sub>-C<sub>5</sub> alkyl substituted with 0-3 R<sup>10</sup>, and C<sub>1</sub>-C<sub>5</sub> alkoxy substituted with 0-3 R<sup>10</sup>, and a bond to S<sub>f</sub>;

R<sup>10</sup> is independently selected at each occurrence from the group: a bond to S<sub>f</sub>, COOR<sup>11</sup>, OH, NHR<sup>11</sup>, C<sub>1-5</sub> alkyl substituted with 0-1 R<sup>12</sup>, and C<sub>1-5</sub> alkoxy substituted with 0-1 R<sup>12</sup>;

R<sup>11</sup> is independently selected at each occurrence from the group: H, aryl substituted with 0-1 R<sup>12</sup>, C<sub>3-10</sub> cycloalkyl substituted with 0-1 R<sup>12</sup>, amino acid substituted with 0-1 R<sup>12</sup>, and a bond to S<sub>f</sub>;

R<sup>12</sup> is a bond to S<sub>f</sub>;

k is selected from 0, 1, and 2;

h is selected from 0, 1, and 2;

h' is selected from 0, 1, 2, 3, 4, and 5;

h'' is selected from 0, 1, 2, 3, 4, and 5;

g is selected from 0, 1, 2, 3, 4, and 5;

g' is selected from 0, 1, 2, 3, 4, and 5;

g'' is selected from 0, 1, 2, 3, 4, and 5;

g''' is selected from 0, 1, 2, 3, 4, and 5;

s is selected from 0, 1, 2, 3, 4, and 5;

s' is selected from 0, 1, 2, 3, 4, and 5;

s'' is selected from 0, 1, 2, 3, 4, and 5;

t is selected from 0, 1, 2, 3, 4, and 5;

t' is selected from 0, 1, 2, 3, 4, and 5;

and a pharmaceutically acceptable salt thereof.

42. (Previously amended) A method according according to Claim 39, wherein the compound is selected from the group:

1-(1,2-Dipalmitoyl-sn-glycero-3-phosphoethanolamino)-12-(cyclo(Arg-Gly-Asp-D-Phe-Lys)-dodecane-1,12-dione SEQ ID NO: 3;

1-(1,2-Dipalmitoyl-sn-glycero-3-phosphoethanolamino)-12-((ω-amino-PEG<sub>3400</sub>-α-carbonyl)-cyclo(Arg-Gly-Asp-D-Phe-Lys))-dodecane-1,12-dione SEQ ID NO: 3; and,

1-(1,2-Dipalmitoyl-sn-glycero-3-phosphoethanolamino)-12-(( $\omega$ -amino-PEG<sub>3400</sub>- $\alpha$ -carbonyl)-Glu-(cyclo(Arg-Gly-Asp-D-Phe-Lys))<sub>2</sub>)-Dodecane-1,12-dione SEQ ID NO: 16.

43. (original) The method according to claim 39, which further comprises a parenterally acceptable and an echogenic gas.

44. (original) The method according to claim 39, further comprising: 1,2-dipalmitoyl-sn-glycero-3-phosphotidic acid, 1,2-dipalmitoyl-sn-glycero-3-phosphatidylcholine, and N-(methoxypolyethylene glycol 5000 carbamoyl)-1,2-dipalmitoyl-sn-glycero-3-phosphatidylethanolamine.

45. (original) The method according to claim 43, wherein, the echogenic gas is a C<sub>2-5</sub> perfluorocarbon.

46. (original) A kit comprising a compound of Claim 39 and a perfusion imaging agent.

47. (original) The method according to claim 1, wherein the vitronectin targeted imaging agent and a perfusion imaging agent have spectrally separable gamma-emission energies.

48. (original) The method according to claim 1, wherein the images are displayed side-by-side to facilitate interpretation of the localization of the vitronectin targeted imaging in the body, relative to the distribution of the perfusion agent in the body.

49. (original) The method according to claim 1, wherein the images are overlaid to facilitate interpretation of the localization of the vitronectin targeted imaging in the body, relative to the distribution of the perfusion agent in the body.

50. (original) The method according to claim 1, for use in concurrent imaging sites of angiogenesis and organ perfusion.

51. (original) The method according to claim 1, for use in diagnosing and localizing sites of angiogenesis and perfusion abnormalities.
52. (original) The method according to claim 1, for use in concurrent detection and localization of sites of endothelial damage and perfusion abnormalities.
53. (original) The method according to claim 1, for use in the concurrent detection and localization of sites of vulnerable plaque and perfusion abnormalities.
54. (original) The method according to claim 1, wherein administering the vitronectin targeted imaging agent and a perfusion imaging agent is concurrent.
55. (original) The method according to claim 1, wherein administering the vitronectin targeted imaging agent and a perfusion imaging agent is sequential.
56. (original) The method according to claim 1, wherein the vitronectin targeted imaging agent and a perfusion imaging agent are administered in a synergistically effective amount.
57. (original) The method according to claim 1, wherein the gamma-emission energies of the vitronectin targeted imaging agent and the perfusion imaging agent are spectrally separable by pulse-height analysis.
58. (original) The method according to claim 1, wherein the difference in gamma emission spectral energies of the vitronectin antagonist diagnostic metallopharmaceutical and the perfusion imaging agent is >10Kev.
59. (original) The method of claim 1 wherein the perfusion imaging agent is a radiolabelled imaging agent, which is radiolabeled with Tc-99m or Tl-201.

60. (original) The method of claim 4 wherein the ultrasound perfusion agent is comprised of a gaseous microbubble or liquid emulsion.

61. (original) The method of claim 4 wherein the ultrasound perfusion agent is a perfluorocarbon gas.

62. (original) The method of claim 4 wherein the ultrasound perfusion agent is a perfluorocarbon liquid.

63. (original) The method of claim 4 wherein the MRI perfusion imaging agent is comprised of Gd(III), Dy(III), Fe(III), or Mn(II).

64. (Canceled).

65. (original) The method of claim 4 wherein the MRI perfusion imaging agent is selected from the group: trisodium (2(R)-((4, 4-diphenylcyclohexy)(hydroxy)phosphoryloxymethyl) diethylenetriaminopentaacetato(6-))-gadolate(3-), gadopentetic acid, gadodiamide, and gadoteridol.

66. (original) The method of claim 4 wherein the MRI perfusion imaging agent is the vitronectin receptor targeted imaging agent which is unbound to the vitronectin receptor.